Docket No.: ARDEN-JACOB-3

Appl. No.: 10/539,790

REMARKS

With regard to the requirement for restriction which is the only point raised

in the Official Action, Applicant hereby provisionally elects to prosecute Group I,

regarding formula I, covering claims 1-11, 14 and 21-24 partially and claim 12,

where Y = oxygen, with traverse, and reserves the right to file a divisional

application or to take such other appropriate measures as deemed necessary to

protect the invention of Group II. Applicant, was also informed that under the

rejoinder rule, group 15 would be eligible for rejoinder if claims in Group I are

deemed allowable.

Applicant believes that the elected preparation claims are so closely related

to the non-elected process claims, that they should remain in the same application

to preserve unity of invention.

The Examiner states that in view of the X-rated references with respect to

Y=O, the restriction is proper. In response, applicant submits that with respect to

the WO00/64988 reference, the ortho-COOH-group in the 1st step is converted into

phosphooxychloride in examples 5 and 16, which is only possible with very robust

dye stuffs of the named compound 9. This process is not generally applicable as

compared to the process claimed here which is based on the use of N-

hydroxysuccinimide -(NHS-) ester.

Furthermore, with respect to WO 02/055512 reference, examples 1, 5 and

14 are shown to utilize NHS esters of fluorescein and naphtafluorescein; however,

no details of the preparation are set forth.

With respect to the EP 167 998 reference, phophorooxychloride is utilized

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for the synthesis, which is only possible in rare cases due to its high reactivity and is not suitable when secondary amines are utilized, which further carry functional groups e.g. COOH. Accordingly the only examples shown are those with simple alkyl-substituents (methyl and ethyl).

Moreover, the special technical feature which all the dye stuffs have in common is the orth-carboxy-phenyl substituents at the central chromophore. The phenyl-group can be additionally substituted or it can also be either a naphthyl, pyridyl or cyclohexyl-group or similar. Essential is only the ortho-COOH-group (generally COOH group which is adjacent to the coupling site of the ring-shaped substituent with the chromophore). Such compounds form in solution an equilibrium between the dye stuff A and a lactone-form B, which in the visible light does not absorb nor fluoresce.

A: Dye (colored)

B: Lactone (colorless)

While COOH groups are actually suitable for coupling top NH₂ groups, for example a protein, this is not the goal here. The coupling takes place but the resulting primary amide immediately forms a lactam, which, similar to the lacton, does not absorb nor fluoresce. The purpose of the fluorescent marking of the biomolecule would thus have no purpose, which is discussed on page 1 of the specificatrion. In the pesent invention, the ortho-COOH group is converted into a secondary amide

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function, so that no lactam can form. A COOH group suitable for coupling or

another function is provided via a hydrocarbon chain.

The instant patent application is based on an international PCT application

pursuant to Chapter II, whereby during the examination of the international patent

application, the unity of the invention was not disputed.

It is believed that the Examiner is trying to draw too fine a line of distinction

and that when all the various facts are taken into account, all claims on file should

be examined on the merits. In any event, Group I and hence claims 1-11, 14 and

21-24 partially and claim 12, where Y = oxygen, are entitled to action on the

merits.

Since applicant has fully and completely responded to the Official Action

and has made the required election, this application is now in order for early action

on the merits of claims

It is noted that each named inventor of the subject matter of the instant

application contributed at least to one of the claims, presently on file.

The Commissioner is hereby authorized to charge any fees which may be

required or credit any overpayment to Deposit Account No.50-1747.

Respectfully submitted

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